

Supplemental Table 1. Complete list of Gene Ontology Biological Processes from the SAFE analysis												
GO term	Neocortex p-values			Heart p-values			Muscle p-values			GO ID	SAFE	
	Age	CR	Resv	Age	CR	Resv	Age	CR	Resv			
autophagic vacuole formation	0.265	0.028	0.006	0.783	0.443	0.013	0.221	0.296	0.227	45		
protein import into nucleus, docking	0.751	0.024	0.095	0.471	0.115	0.516	0.505	0.026	0.110	59		
protein import into nucleus, translocation	0.774	0.177	0.261	0.336	0.444	0.114	0.617	0.025	0.046	60		
regulation of progression through cell cycle	0.969	0.015	0.002	0.073	0.040	0.004	0.259	0.082	0.328	74		
DNA damage checkpoint	0.847	0.861	0.710	0.264	0.279	0.287	0.252	0.882	1.000	77		
regulation of cyclin-dependent protein kinase activity	0.486	0.146	0.365	0.123	0.659	0.019	0.054	0.799	0.420	79		
G1/S transition of mitotic cell cycle	0.679	0.207	0.387	0.650	0.010	0.852	0.993	0.142	0.538	82		
G2/M transition of mitotic cell cycle	0.944	0.229	0.141	0.378	0.829	0.183	0.047	0.890	0.648	86		
negative regulation of transcription from RNA polymerase II promoter	0.377	0.051	0.006	0.644	0.030	0.034	0.473	0.560	0.540	122		
two-component signal transduction system (phosphorelay)	0.020	0.081	0.199	0.449	0.370	0.542	0.164	0.366	0.645	160		
MAPKK cascade	0.472	0.066	0.591	0.081	0.055	0.396	0.006	0.060	0.119	165		
mRNA catabolic process, nonsense-mediated decay	0.232	0.087	0.006	0.135	0.078	0.033	0.238	0.292	0.003	184		
activation of MAPK activity	0.468	0.704	0.958	0.440	0.482	0.380	0.075	0.441	0.397	187		
inactivation of MAPK activity	0.016	0.064	0.019	0.078	0.107	0.290	0.620	0.612	0.792	188		
microtubule cytoskeleton organization and biogenesis	0.469	0.688	0.630	0.056	0.531	0.005	0.037	0.848	0.297	226		
spliceosome assembly	0.924	0.018	0.176	0.045	0.661	0.519	0.524	0.508	0.049	245		
peptidoglycan metabolic process	0.538	0.876	0.583	0.852	0.835	0.904	0.395	0.706	0.934	270		
nuclear mRNA splicing, via spliceosome	0.922	0.010	0.006	0.023	0.015	0.032	0.159	0.078	0.003	396		
telomere maintenance	0.868	0.725	0.121	0.053	0.707	0.808	0.827	0.411	0.520	723		
cell morphogenesis	0.520	0.757	0.806	0.066	0.120	0.141	0.819	0.677	0.400	902		
cytokinesis	0.043	0.888	0.115	0.539	0.169	0.585	0.759	0.367	0.169	910		
skeletal development	0.639	0.663	0.748	0.576	0.465	0.744	0.108	0.540	0.740	1501		
cartilage condensation	0.062	0.128	0.096	0.617	0.234	0.193	0.036	0.164	0.374	1502		
ossification	0.730	0.520	0.733	0.092	0.617	0.214	0.046	0.006	0.241	1503		
angiogenesis	0.016	0.048	0.146	0.043	0.007	0.030	0.034	0.337	0.042	1525		
ovarian follicle development	0.454	0.241	0.416	0.860	0.708	0.781	0.729	0.926	0.692	1541		
regulation of cell growth	0.665	0.237	0.150	0.008	0.015	0.012	0.152	0.038	0.003	1558		
blood vessel development	0.265	0.369	0.387	0.169	0.045	0.293	0.424	0.060	0.012	1568		
patterning of blood vessels	0.279	0.158	0.811	0.034	0.442	0.125	0.721	0.378	0.131	1569		
vasculogenesis	0.180	0.198	0.605	0.038	0.092	0.706	0.516	0.309	0.305	1570		
osteoblast differentiation	0.240	0.567	0.724	0.768	0.453	0.159	0.210	0.006	0.012	1649		
metanephros development	0.704	0.393	0.715	0.534	0.770	0.693	0.279	0.378	0.629	1656		
ureteric bud development	0.617	0.165	0.659	0.584	0.496	0.733	0.141	0.088	0.309	1657		
ureteric bud branching	0.200	0.809	0.341	0.527	0.640	0.907	0.696	0.378	0.675	1658		
response to hypoxia	0.361	0.054	0.055	0.007	0.067	0.005	0.658	0.701	0.734	1666		
in utero embryonic development	0.960	0.278	0.030	0.007	0.047	0.005	0.232	0.299	0.570	1701		
mesoderm formation	0.163	0.111	0.028	0.285	0.070	0.005	0.790	0.006	0.014	1707		
cell fate specification	0.080	0.127	0.858	0.433	0.341	0.494	0.526	0.550	0.952	1708		
cell fate determination	0.397	0.190	0.432	0.135	0.025	0.064	0.492	0.909	0.804	1709		
eye development (sensu Mammalia)	0.262	0.122	0.233	0.512	0.520	0.213	0.164	0.042	0.705	1747		
neural crest cell migration	0.297	0.050	0.196	0.587	0.571	0.725	0.550	0.020	0.103	1755		
somitogenesis	0.922	0.557	0.943	0.080	0.359	0.167	0.876	0.879	0.919	1756		
induction of an organ	0.868	0.428	0.928	0.888	0.831	0.950	0.492	0.560	1.000	1759		
neuron migration	0.115	0.289	0.320	0.065	0.953	0.731	0.151	0.017	0.307	1764		
cytokine production	0.600	0.942	0.503	0.324	0.276	0.037	0.425	0.074	0.436	1816		
kidney development	0.633	0.609	0.977	0.723	0.952	0.916	0.928	0.486	0.979	1822		
neural tube closure	0.094	0.302	0.090	0.189	0.167	0.592	0.503	0.371	0.462	1843		
placenta development	0.984	0.610	0.718	0.837	0.621	0.775	0.982	0.749	0.509	1890		
heart looping	0.990	0.368	0.223	0.029	0.122	0.305	0.577	0.617	0.591	1947		
blood vessel remodeling	0.260	0.404	0.193	0.674	0.341	0.211	0.984	0.940	0.831	1974		
morphogenesis of an epithelium	0.776	0.499	0.718	0.846	0.683	0.896	0.289	0.388	0.549	2009		
antigen processing and presentation of peptide antigen via MHC	0.127	0.937	0.332	0.992	0.874	0.948	0.178	0.815	0.628	2474		
carbohydrate metabolic process	0.760	0.321	0.035	0.008	0.035	0.030	0.046	0.752	0.690	5975		
glycogen metabolic process	0.943	0.315	0.033	0.008	0.191	0.062	0.110	0.120	0.412	5977		
glucose metabolic process	0.323	0.309	0.925	0.016	0.032	0.021	0.008	0.601	0.766	6006		
N-acetylglucosamine metabolic process	0.579	0.418	0.473	0.070	0.582	0.135	0.559	0.903	0.817	6044		
glycerol metabolic process	0.797	0.100	0.814	0.027	0.007	0.041	0.208	0.637	0.262	6071		
generation of precursor metabolites and energy	0.923	0.287	0.352	0.706	0.376	0.354	0.097	0.591	0.404	6091		
gluconeogenesis	0.680	0.563	0.861	0.505	0.070	0.065	0.175	0.027	0.324	6094		
glycolysis	0.858	0.033	0.379	0.642	0.068	0.054	0.413	0.375	0.035	6096		
tricarboxylic acid cycle	0.749	0.043	0.126	0.310	0.038	0.028	0.042	0.846	0.003	6099		
electron transport	0.450	0.655	0.266	0.299	0.103	0.154	0.077	0.381	0.053	6118		
mitochondrial electron transport, NADH to ubiquinone	0.192	0.043	0.325	0.266	0.301	0.246	0.430	0.523	0.014	6120		
nucleobase, nucleoside, nucleotide and nucleic acid metabolic process	0.854	0.150	0.616	0.018	0.593	0.310	0.267	0.429	0.333	6139		
purine nucleotide biosynthetic process	0.766	0.207	0.052	0.047	0.030	0.033	0.137	0.072	0.015	6164		
cAMP biosynthetic process	0.251	0.095	0.742	0.067	0.595	0.418	0.337	0.704	0.773	6171		
cGMP biosynthetic process	0.561	0.853	0.341	0.276	0.233	0.670	0.583	0.253	0.115	6182		
GTP biosynthetic process	0.732	0.348	0.184	0.289	0.933	0.519	0.903	0.659	0.360	6183		
UTP biosynthetic process	0.629	0.528	0.240	0.260	0.927	0.432	0.976	0.793	0.572	6228		
CTP biosynthetic process	0.798	0.637	0.261	0.350	0.893	0.470	0.912	0.700	0.380	6241		
DNA metabolic process	0.253	0.680	0.183	0.444	0.697	0.542	0.074	0.512	0.779	6259		
DNA replication	0.261	0.160	0.013	0.057	0.007	0.013	0.625	0.632	0.340	6260		
DNA-dependent DNA replication	0.379	0.420	0.833	0.451	0.122	0.537	0.266	0.312	0.146	6261		
DNA topological change	0.708	0.583	0.563	0.028	0.119	0.347	0.778	0.233	0.135	6265		
DNA unwinding during replication	0.757	0.810	0.518	0.364	0.474	0.215	0.828	0.205	0.413	6268		
DNA replication initiation	0.104	0.566	0.273	0.247	0.424	0.322	1.000	0.395	0.439	6270		
DNA repair	0.585	0.222	0.068	0.008	0.010	0.005	0.002	0.796	0.646	6281		
base-excision repair	0.586	0.176	0.021	0.052	0.341	0.347	0.512	0.769	0.197	6284		
nucleotide-excision repair	0.754	0.262	0.236	0.058	0.180	0.603	0.462	0.970	0.265	6289		
mismatch repair	0.772	0.855	0.271	0.324	0.475	0.713	0.666	0.895	0.954	6298		
double-strand break repair	0.248	0.360	0.214	0.259	0.718	0.129	0.402	0.775	0.660	6302		
DNA methylation	0.692	0.306	0.773	0.059	0.581	0.177	0.601	0.438	0.328	6306		
DNA fragmentation during apoptosis	0.289	0.435	0.779	0.062	0.118	0.384	0.626	0.669	0.794	6309		
DNA recombination	0.757	0.477	0.094	0.075	0.294	0.129	0.049	0.088	0.753	6310		
DNA packaging	0.833	0.017	0.103	0.257	0.015	0.052	0.843	0.134	0.065	6323		
establishment and/or maintenance of chromatin architecture	0.310	0.099	0.056	0.703	0.086	0.753	0.921	0.308	0.065	6325		
chromatin assembly or disassembly	0.310	0.022	0.006	0.033	0.036	0.027	0.161	0.016	0.045	6333		
nucleosome assembly	0.977	0.184	0.282	0.402	0.187	0.172	0.572	0.392	0.091	6334		
chromatin remodeling	0.433	0.144	0.045	0.178	0.035	0.069	0.924	0.133	0.209	6338		
chromatin silencing	0.807	0.230	0.208	0.544	0.455	0.655	0.595	0.125	0.399	6342		
transcription	0.253	0.005	0.002	0.043	0.012	0.003	0.482	0.134	0.020	6350		

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GO term	Neocortex p-values			Heart p-values			Muscle p-values			GO ID	SAFE	
	Age	CR	Resv	Age	CR	Resv	Age	CR	Resv			
mRNA polyadenylation	0.955	0.746	0.518	0.330	0.514	0.339	0.961	0.530	0.442	6378		
RNA processing	0.653	0.010	0.016	0.095	0.141	0.070	0.621	0.078	0.120	6396		
mRNA processing	0.634	0.004	0.006	0.006	0.036	0.005	0.277	0.127	0.025	6397		
RNA catabolic process	0.871	0.079	0.524	0.025	0.111	0.015	0.891	0.224	0.121	6401		
mRNA catabolic process	0.424	0.231	0.541	0.024	0.057	0.017	0.397	0.297	0.138	6402		
mRNA export from nucleus	0.828	0.235	0.018	0.046	0.137	0.206	0.094	0.905	0.070	6406		
translation	0.854	0.008	0.047	0.008	0.010	0.005	0.253	0.498	0.003	6412		
translational initiation	0.553	0.010	0.005	0.006	0.007	0.005	0.277	0.641	0.009	6413		
translational elongation	0.862	0.067	0.006	0.023	0.089	0.033	0.037	0.630	0.014	6414		
regulation of translation	0.143	0.134	0.034	0.172	0.317	0.023	0.410	0.172	0.039	6417		
tRNA aminoacylation for protein translation	0.740	0.038	0.014	0.064	0.045	0.246	0.250	0.077	0.022	6418		
regulation of protein biosynthesis DUPLICATE	0.556	0.113	0.046	0.034	0.516	0.015	0.110	0.545	0.025	6445		
regulation of translational initiation	0.500	0.079	0.706	0.025	0.069	0.085	0.142	0.091	0.009	6446		
protein folding	0.306	0.007	0.026	0.036	0.007	0.013	0.266	0.232	0.344	6457		
protein complex assembly	0.319	0.010	0.003	0.046	0.467	0.164	0.201	0.246	0.061	6461		
protein modification process	0.068	0.007	0.005	0.008	0.061	0.005	0.407	0.019	0.016	6464		
protein amino acid phosphorylation	0.203	0.031	0.006	0.016	0.007	0.024	0.184	0.084	0.041	6468		
negative regulation of protein kinase activity	0.017	0.292	0.034	0.008	0.665	0.056	0.234	0.934	0.941	6469		
protein amino acid dephosphorylation	0.519	0.022	0.028	0.027	0.007	0.005	0.226	0.123	0.019	6470		
protein amino acid ADP-ribosylation	0.595	0.661	0.019	0.095	0.112	0.190	0.359	0.006	0.044	6471		
protein amino acid methylation	0.048	0.172	0.756	0.106	0.191	0.196	0.499	0.047	0.194	6479		
protein amino acid glycosylation	0.487	0.313	0.028	0.405	0.374	0.219	0.021	0.583	0.670	6486		
protein amino acid N-linked glycosylation	0.815	0.019	0.026	0.651	0.285	0.743	0.810	0.032	0.632	6487		
protein amino acid O-linked glycosylation	0.271	0.066	0.301	0.376	0.142	0.874	0.483	0.044	0.043	6493		
GPI anchor biosynthetic process	0.985	0.187	0.198	0.280	0.656	0.742	0.379	0.975	0.161	6506		
proteolysis	0.069	0.626	0.871	0.285	0.174	0.111	0.422	0.778	0.953	6508		
membrane protein ectodomain proteolysis	0.008	0.302	0.298	0.165	0.230	0.104	0.834	0.716	0.176	6509		
ubiquitin-dependent protein catabolic process	0.332	0.010	0.006	0.034	0.007	0.019	0.408	0.210	0.018	6511		
ubiquitin cycle	0.505	0.010	0.006	0.031	0.007	0.012	0.387	0.019	0.003	6512		
amino acid metabolic process	0.439	0.289	0.294	0.512	0.007	0.091	0.044	0.680	0.170	6520		
glutamine metabolic process	0.701	0.284	0.295	0.149	0.638	0.316	0.255	0.047	0.019	6541		
protein targeting	0.406	0.225	0.133	0.016	0.057	0.087	0.150	0.180	0.025	6605		
protein import into nucleus	0.479	0.206	0.445	0.445	0.094	0.444	0.239	0.047	0.012	6606		
protein export from nucleus	0.378	0.025	0.101	0.650	0.380	0.450	0.969	0.493	0.361	6611		
protein targeting to mitochondrion	0.489	0.062	0.068	0.230	0.051	0.008	0.402	0.320	0.219	6626		
lipid metabolic process	0.034	0.040	0.432	0.118	0.023	0.061	0.584	0.225	0.466	6629		
fatty acid metabolic process	0.474	0.362	0.082	0.632	0.043	0.053	0.392	0.323	0.030	6631		
fatty acid biosynthetic process	0.215	0.688	0.729	0.110	0.097	0.194	0.306	0.105	0.198	6633		
fatty acid beta-oxidation	0.324	0.350	0.348	0.033	0.039	0.027	0.070	0.268	0.012	6635		
acyl-CoA metabolic process	0.340	0.382	0.500	0.183	0.459	0.055	0.770	0.902	0.954	6637		
phospholipid metabolic process	0.520	0.968	0.583	0.373	0.321	0.798	0.295	0.142	0.824	6644		
sphingolipid metabolic process	0.956	0.358	0.840	0.185	0.076	0.034	0.545	0.255	0.748	6665		
ceramide metabolic process	0.579	0.063	0.075	0.911	0.908	0.355	0.017	0.570	0.527	6672		
leukotriene metabolic process	0.635	0.920	0.272	0.356	0.769	0.743	0.953	0.616	0.650	6691		
steroid biosynthetic process	0.083	0.344	0.959	0.074	0.387	0.127	0.071	0.417	0.353	6694		
cholesterol biosynthetic process	0.457	0.147	0.794	0.114	0.027	0.083	0.125	0.294	0.320	6695		
C21-steroid hormone biosynthetic process	0.223	0.430	0.971	0.973	0.916	0.448	0.771	0.900	0.944	6700		
aromatic compound metabolic process	0.018	0.060	0.031	0.659	0.091	0.106	0.088	0.768	0.425	6725		
one-carbon compound metabolic process	0.018	0.501	0.775	0.049	0.482	0.569	0.143	0.632	0.545	6730		
glutathione metabolic process	0.018	0.432	0.252	0.189	0.007	0.082	0.064	0.699	0.062	6749		
ATP biosynthetic process	0.602	0.043	0.464	0.438	0.148	0.187	0.042	0.211	0.389	6754		
heme biosynthetic process	0.313	0.730	0.391	0.161	0.007	0.046	0.275	0.626	0.546	6783		
sulfur metabolic process	0.497	0.833	0.619	0.301	0.840	0.518	0.619	0.623	0.684	6790		
phosphate metabolic process	0.917	0.196	0.113	0.046	0.760	0.182	0.254	0.571	0.606	6796		
superoxide metabolic process	0.139	0.373	0.460	0.074	0.039	0.838	0.424	0.278	0.366	6801		
xenobiotic metabolic process	0.066	0.670	0.425	0.436	0.867	0.146	0.285	0.717	0.868	6805		
nitrogen compound metabolic process	0.759	0.031	0.078	0.715	0.083	0.181	0.771	0.083	0.096	6807		
nitric oxide biosynthetic process	0.622	0.347	0.153	0.784	0.082	0.541	0.901	0.372	0.870	6809		
transport	0.085	0.004	0.004	0.006	0.007	0.004	0.307	0.410	0.111	6810		
ion transport	0.020	0.383	0.233	0.983	0.788	0.953	0.807	0.471	0.905	6811		
cation transport	0.117	0.762	0.090	0.903	0.492	0.579	0.577	0.706	0.194	6812		
potassium ion transport	0.562	0.658	0.187	0.960	0.535	0.452	0.579	0.233	0.803	6813		
sodium ion transport	0.082	0.502	0.228	0.537	0.569	0.965	0.622	0.731	0.705	6814		
calcium ion transport	0.057	0.147	0.031	0.955	0.116	0.655	0.683	0.612	0.909	6816		
phosphate transport	0.970	0.366	0.787	0.250	0.136	0.384	0.419	0.820	0.808	6817		
hydrogen transport	0.923	0.468	0.254	0.741	0.813	0.470	0.289	0.619	0.270	6818		
anion transport	0.262	0.177	0.120	0.537	0.098	0.233	0.689	0.148	0.175	6820		
chloride transport	0.680	0.810	0.319	0.513	0.543	0.429	1.000	0.859	0.647	6821		
iron ion transport	0.284	0.577	0.474	0.008	0.112	0.023	0.055	0.040	0.128	6826		
zinc ion transport	0.651	0.599	0.398	0.098	0.586	0.659	0.672	0.458	0.257	6829		
water transport	0.541	0.944	0.944	0.800	0.417	0.576	0.779	0.200	0.479	6833		
neurotransmitter transport	0.855	0.147	0.853	0.661	0.826	0.916	0.930	0.186	0.922	6836		
mitochondrial transport	0.311	0.137	0.064	0.042	0.491	0.111	0.087	0.650	0.333	6839		
amino acid transport	0.762	0.185	0.117	0.431	0.899	0.951	0.901	0.237	0.801	6865		
lipid transport	0.006	0.042	0.025	0.095	0.023	0.021	0.730	0.251	0.493	6869		
calcium ion homeostasis	0.030	0.553	0.737	0.679	0.539	0.408	0.104	0.615	0.710	6874		
iron ion homeostasis	0.757	0.779	0.516	0.008	0.073	0.015	0.006	0.019	0.357	6879		
regulation of pH	0.436	0.197	0.136	0.057	0.537	0.990	0.558	0.546	0.466	6885		
intracellular protein transport	0.452	0.010	0.005	0.151	0.037	0.019	0.382	0.134	0.011	6886		
exocytosis	0.136	0.029	0.016	0.412	0.232	0.420	0.742	0.301	0.322	6887		
ER to Golgi vesicle-mediated transport	0.311	0.010	0.006	0.014	0.020	0.044	0.508	0.386	0.056	6888		
intra-Golgi vesicle-mediated transport	0.667	0.062	0.087	0.270	0.057	0.033	0.343	0.128	0.360	6891		
endocytosis	0.587	0.026	0.046	0.155	0.163	0.005	0.222	0.027	0.002	6897		
receptor-mediated endocytosis	0.891	0.789	0.183	0.603	0.632	0.163	0.068	0.024	0.463	6898		
vesicle docking during exocytosis	0.618	0.018	0.068	0.500	0.678	0.533	0.575	0.801	0.779	6904		
phagocytosis	0.203	0.178	0.039	0.591	0.784	0.886	0.694	0.845	0.960	6909		
phagocytosis, recognition	0.140	0.155	0.424	0.346	0.055	0.179	0.021	0.018	0.088	6910		
phagocytosis, engulfment	0.018	0.488	0.649	0.350	0.062	0.474	0.016	0.010	0.109	6911		
nucleocytoplasmic transport	0.083	0.241	0.016	0.583	0.135	0.147	0.368	0.705	0.166	6913		
autophagy	0.803	0.048	0.054	0.587	0.195	0.024	0.412	0.069	0.029	6914		
apoptosis	0.015	0.022	0.004	0.008	0.048	0.005	0.499	0.077				

Supplemental Table 1. Complete list of Gene Ontology Biological Processes from the SAFE analysis													
GO term	Neocortex p-values			Heart p-values			Muscle p-values			GO ID	SAFE		
	Age	CR	Resv	Age	CR	Resv	Age	CR	Resv				
smooth muscle contraction	0.386	0.351	0.985	0.079	0.472	0.068	0.144	0.620	0.588	6939			
striated muscle contraction	0.780	0.943	0.938	0.460	0.056	0.307	0.083	0.832	0.135	6941			
membrane fusion	0.760	0.134	0.006	0.932	0.357	0.033	0.254	0.193	0.561	6944			
response to stress	0.443	0.140	0.340	0.008	0.033	0.040	0.143	0.373	0.213	6950			
defense response	0.948	0.991	0.992	0.999	0.990	0.948	0.953	0.962	0.970	6952			
acute-phase response	0.785	0.898	0.863	0.684	0.746	0.764	0.823	0.968	0.669	6953			
inflammatory response	0.047	0.986	0.994	0.693	0.825	0.845	0.248	0.304	0.285	6954			
immune response	0.226	0.943	0.995	1.000	0.972	0.967	0.234	0.461	0.778	6955			
complement activation	0.120	0.314	0.871	0.926	0.523	0.324	0.306	0.769	0.585	6956			
complement activation, alternative pathway	0.788	0.579	0.935	0.461	0.596	0.814	0.471	0.916	0.506	6957			
complement activation, classical pathway	0.356	0.659	0.997	0.919	0.697	0.292	0.075	0.795	0.769	6958			
humoral immune response	0.537	0.796	0.882	0.946	0.921	0.811	0.234	0.953	0.708	6959			
cellular defense response	0.525	0.967	0.972	0.565	0.520	0.760	0.620	0.850	0.836	6966			
response to DNA damage stimulus	0.605	0.464	0.194	0.008	0.020	0.038	0.002	0.312	0.303	6974			
response to oxidative stress	0.661	0.076	0.166	0.008	0.007	0.094	0.034	0.238	0.340	6979			
response to unfolded protein	0.593	0.025	0.209	0.017	0.049	0.013	0.130	0.039	0.684	6986			
chromosome organization and biogenesis (sensu Eukaryota)	0.569	0.450	0.322	0.118	0.246	0.350	0.756	0.187	0.234	7001			
telomere maintenance via telomerase	0.104	0.534	0.696	0.066	0.375	0.597	0.364	0.842	0.430	7004			
cytoskeleton organization and biogenesis	0.695	0.588	0.962	0.452	0.013	0.038	0.437	0.781	0.195	7010			
actin filament organization	0.085	0.018	0.042	0.030	0.184	0.435	0.293	0.016	0.015	7015			
microtubule-based process	0.138	0.052	0.006	0.068	0.033	0.143	0.035	0.589	0.777	7017			
microtubule-based movement	0.451	0.089	0.006	0.491	0.653	0.901	0.579	0.605	0.900	7018			
peroxisome organization and biogenesis	0.138	0.010	0.006	0.152	0.026	0.015	0.531	0.871	0.014	7031			
lysosome organization and biogenesis	0.076	0.157	0.062	0.245	0.649	0.314	0.083	0.570	0.116	7040			
ribosome biogenesis	0.845	0.010	0.110	0.160	0.051	0.025	0.020	0.594	0.243	7046			
cell cycle	0.499	0.022	0.005	0.124	0.007	0.017	0.272	0.456	0.058	7049			
cell cycle arrest	0.899	0.110	0.058	0.028	0.039	0.046	0.196	0.236	0.157	7050			
chromosome segregation	0.099	0.539	0.084	0.028	0.288	0.197	0.342	0.230	0.353	7059			
mitosis	0.269	0.113	0.006	0.056	0.013	0.005	0.243	0.657	0.400	7067			
mitotic chromosome condensation	0.402	0.097	0.208	0.839	0.392	0.032	0.433	0.970	0.941	7076			
meiosis	0.716	0.592	0.681	0.783	0.412	0.977	0.347	0.873	0.987	7126			
meiotic recombination	0.436	0.894	0.788	0.454	0.615	0.398	0.271	0.683	0.843	7131			
cell communication	0.267	0.017	0.102	0.096	0.461	0.176	0.327	0.508	0.201	7154			
cell adhesion	0.005	0.064	0.024	0.288	0.067	0.503	0.055	0.440	0.661	7155			
homophilic cell adhesion	0.131	0.695	0.274	0.781	0.466	0.630	0.371	0.889	1.000	7156			
heterophilic cell adhesion	0.254	0.047	0.026	0.674	0.393	0.777	0.018	0.360	0.638	7157			
cell-matrix adhesion	0.045	0.587	0.615	0.027	0.033	0.550	0.126	0.325	0.755	7160			
negative regulation of cell adhesion	0.129	0.233	0.395	0.746	0.152	0.057	0.249	0.629	0.686	7162			
establishment and/or maintenance of cell polarity	0.600	0.060	0.095	0.038	0.287	0.039	0.381	0.023	0.265	7163			
signal transduction	0.145	0.680	0.949	0.767	0.491	0.304	0.044	0.012	0.803	7165			
cell surface receptor linked signal transduction	0.658	0.951	0.975	0.992	0.845	0.723	0.558	0.380	0.712	7166			
transmembrane receptor protein tyrosine kinase signaling pathway	0.301	0.752	0.846	0.532	0.755	0.463	0.486	0.275	0.682	7169			
epidermal growth factor receptor signaling pathway	0.140	0.262	0.125	0.214	0.056	0.128	0.793	0.070	0.114	7173			
transmembrane receptor protein serine/threonine kinase signaling	0.577	0.707	0.587	0.180	0.088	0.050	0.145	0.015	0.160	7178			
transforming growth factor beta receptor signaling pathway	0.422	0.549	0.539	0.516	0.504	0.361	0.206	0.111	0.449	7179			
transmembrane receptor protein tyrosine phosphatase signaling	0.136	0.660	0.773	0.482	0.299	0.727	0.797	0.442	0.464	7185			
G-protein coupled receptor protein signaling pathway	0.128	0.980	1.000	0.980	0.999	0.998	0.555	0.379	0.962	7186			
G-protein signaling, coupled to cyclic nucleotide second messenger	0.885	0.784	0.337	0.834	0.885	0.752	0.217	0.370	0.790	7187			
G-protein signaling, coupled to cAMP nucleotide second messenger	0.278	0.169	0.987	0.197	0.155	0.761	0.567	0.627	0.884	7188			
G-protein signaling, adenylyl cyclase activating pathway	0.533	0.413	0.436	0.864	0.649	0.343	0.040	0.413	0.118	7189			
adenylyl cyclase activation	0.803	0.334	0.935	0.205	0.937	0.908	0.192	0.121	0.071	7190			
G-protein signaling, adenylyl cyclase inhibiting pathway	0.313	0.193	0.006	0.548	0.026	0.367	0.080	0.216	0.979	7193			
G-protein signaling, coupled to IP3 second messenger (phosphoinositide cycle)	0.121	0.883	0.968	0.610	0.382	0.987	0.717	0.522	0.396	7200			
phospholipase C activation	0.742	0.432	0.753	0.125	0.229	0.289	0.696	0.170	0.614	7202			
elevation of cytosolic calcium ion concentration	0.398	0.514	0.997	0.941	0.755	0.989	0.418	0.707	0.862	7204			
protein kinase C activation	0.399	0.019	0.233	0.534	0.136	0.308	0.546	0.850	0.751	7205			
gamma-aminobutyric acid signaling pathway	0.373	0.676	0.066	0.949	0.577	0.621	0.932	0.443	0.158	7214			
glutamate signaling pathway	0.419	0.088	0.064	0.924	0.287	0.528	0.351	0.245	0.594	7215			
neuropeptide signaling pathway	0.051	0.357	0.672	0.743	0.708	0.912	0.445	0.407	0.996	7218			
Notch signaling pathway	0.443	0.035	0.278	0.058	0.129	0.013	0.132	0.006	0.024	7219			
frizzled signaling pathway	0.924	0.054	0.118	0.288	0.069	0.487	0.006	0.054	0.009	7222			
Wnt receptor signaling pathway, calcium modulating pathway	0.817	0.812	0.978	0.772	0.695	0.801	0.811	0.845	0.972	7223			
smoothened signaling pathway	0.376	0.700	0.763	0.379	0.131	0.379	0.769	0.119	0.490	7224			
integrin-mediated signaling pathway	0.961	0.377	0.780	0.177	0.282	0.532	0.479	0.781	0.999	7229			
intracellular signaling cascade	0.096	0.005	0.077	0.001	0.046	0.067	0.267	0.089	0.101	7242			
protein kinase cascade	0.226	0.010	0.062	0.053	0.147	0.067	0.158	0.714	0.122	7243			
I-kappaB kinase/NF-kappaB cascade	0.178	0.934	0.952	0.618	0.890	0.330	0.832	0.065	0.226	7249			
JNK cascade	0.154	0.010	0.437	0.524	0.110	0.047	0.481	0.218	0.655	7254			
activation of JNK activity	0.099	0.325	0.853	0.345	0.022	0.074	0.646	0.138	0.248	7257			
JAK-STAT cascade	0.548	0.342	0.179	0.054	0.221	0.293	0.851	0.300	0.031	7259			
small GTPase mediated signal transduction	0.034	0.004	0.002	0.002	0.007	0.005	0.614	0.258	0.058	7264			
Ras protein signal transduction	0.498	0.265	0.092	0.224	0.161	0.030	0.106	0.328	0.328	7265			
Rho protein signal transduction	0.202	0.018	0.016	0.008	0.024	0.005	0.761	0.426	0.067	7266			
cell-cell signaling	0.332	0.822	0.997	0.903	0.962	0.653	0.343	0.161	0.997	7267			
synaptic transmission	0.164	0.052	0.028	0.956	0.956	0.862	0.478	0.640	0.911	7268			
neurotransmitter secretion	0.297	0.202	0.006	0.908	0.807	0.881	0.209	0.029	0.073	7269			
synaptic transmission, cholinergic	0.376	0.766	0.823	0.086	0.903	0.202	0.633	0.361	0.404	7271			
multicellular organismal development	0.504	0.275	0.996	0.442	0.837	0.586	0.102	0.289	0.739	7275			
gamete generation	0.063	0.263	0.675	0.530	0.044	0.784	0.101	0.104	0.683	7276			
germ cell development	0.235	0.010	0.287	0.649	0.131	0.272	0.819	0.915	0.711	7281			
spermatogenesis	0.688	0.956	0.927	0.959	0.865	0.632	0.752	0.637	0.938	7283			
spermatid development	0.752	0.109	0.619	0.387	0.510	0.354	0.124	0.213	0.003	7286			
female gamete generation	0.967	0.946	0.890	0.383	0.820	0.409	0.050	0.436	0.998	7292			
single fertilization	0.994	0.748	0.258	0.125	0.683	0.440	0.569	0.678	0.676	7338			
binding of sperm to zona pellucida	0.778	0.764	0.713	1.000	0.877	0.979	0.772	0.480	0.944	7339			
acrosome reaction	0.516	0.600	0.829	0.337	0.400	0.449	0.791	0.067	0.803	7340			
determination of left/right symmetry	0.496	0.558	0.987	0.217	0.159	0.537	0.300	0.141	0.407	7368			
pattern specification process	0.928	0.971	0.958	0.964	0.558	0.925	0.745	0.459	0.520	7389			
ectoderm development	0.338	0.517	0.591	0.742	0.776	0.480	0.466	0.831	0.972	7398			
nervous system development	0.058	0.010	0.005	0.114	0.941	0.665	0.021	0.412	0.956	7399			
axonogenesis	0.045	0.144</											

Supplemental Table 1. Complete list of Gene Ontology Biological Processes from the SAFE analysis

GO term	Neocortex p-values			Heart p-values			Muscle p-values			GO ID	SAFE
	Age	CR	Resv	Age	CR	Resv	Age	CR	Resv		
mesoderm development	0.818	0.649	0.666	0.167	0.201	0.070	0.872	0.442	0.891	7498	
heart development	0.555	0.096	0.274	0.218	0.007	0.017	0.317	0.138	0.479	7507	
muscle development	0.943	0.960	0.996	0.056	0.007	0.169	0.377	0.511	0.089	7517	
striated muscle development	0.994	0.256	0.768	0.059	0.087	0.502	0.227	0.121	0.039	7519	
sex differentiation	0.978	0.929	0.980	0.523	0.883	0.867	0.818	0.362	0.994	7548	
pregnancy	0.712	0.851	0.988	0.392	0.283	0.546	0.549	0.225	0.037	7565	
embryo implantation	0.124	0.734	0.316	0.715	0.176	0.460	0.888	0.345	0.826	7566	
cell aging	0.167	0.236	0.058	0.554	0.730	0.672	0.763	0.666	0.898	7569	
physiological process	0.436	0.469	0.201	0.754	0.123	0.370	0.959	0.884	0.750	7582	
response to nutrient	0.015	0.061	0.190	0.068	0.058	0.319	0.129	0.165	0.454	7584	
respiratory gaseous exchange	0.421	0.993	0.989	0.211	0.465	0.199	0.703	0.117	0.476	7585	
digestion	0.394	0.775	0.946	0.950	0.597	0.848	0.289	0.910	0.913	7586	
excretion	0.787	0.913	0.991	0.789	0.180	0.253	0.278	0.191	0.927	7588	
lactation	0.488	0.025	0.232	0.162	0.919	0.532	0.094	0.657	0.308	7595	
blood coagulation	0.057	0.603	0.807	0.784	0.284	0.187	0.310	0.298	0.406	7596	
sensory perception	0.238	0.027	0.476	0.958	0.915	0.930	0.168	0.482	0.340	7600	
visual perception	0.941	0.346	0.265	0.923	0.880	0.816	0.097	0.887	0.984	7601	
phototransduction	0.898	0.759	0.355	0.291	0.768	0.807	0.146	0.946	0.766	7602	
sensory perception of sound	0.383	0.343	0.649	0.829	0.620	0.963	0.847	0.273	0.388	7605	
sensory perception of chemical stimulus	0.947	0.887	0.990	0.872	0.821	0.706	0.204	0.031	0.075	7606	
sensory perception of smell	0.709	0.983	1.000	0.851	0.664	0.976	0.600	0.897	0.926	7608	
behavior	0.175	0.178	0.272	0.053	0.663	0.657	0.077	0.345	0.886	7610	
learning and/or memory	0.036	0.269	0.388	0.772	0.494	0.368	0.300	0.379	0.703	7611	
learning	0.170	0.164	0.171	0.943	0.669	0.301	0.071	0.006	0.361	7612	
memory	0.205	0.101	0.021	0.019	0.020	0.012	0.800	0.086	0.881	7613	
circadian rhythm	0.064	0.708	0.083	0.193	0.303	0.734	0.791	0.573	0.952	7623	
locomotory behavior	0.064	0.128	0.006	0.718	0.620	0.789	0.635	0.825	0.917	7626	
adult walking behavior	0.474	0.757	0.077	0.195	0.242	0.443	0.155	0.488	0.897	7628	
feeding behavior	0.072	0.786	0.962	0.713	0.420	0.960	0.387	0.908	0.922	7631	
circulation	0.008	0.420	0.729	0.608	0.072	0.082	0.014	0.099	0.044	8015	
regulation of heart contraction	0.365	0.036	0.043	0.436	0.256	0.060	0.093	0.816	0.908	8016	
tRNA processing	0.980	0.267	0.513	0.008	0.558	0.090	0.867	0.259	0.014	8033	
protein localization	0.075	0.104	0.230	0.156	0.669	0.456	0.489	0.017	0.076	8104	
biological process	0.004	0.008	0.001	0.034	0.023	0.003	0.299	0.528	0.067	8150	
metabolic process	0.023	0.269	0.013	0.035	0.033	0.012	0.101	0.936	0.033	8152	
steroid metabolic process	0.069	0.061	0.323	0.098	0.135	0.533	0.414	0.274	0.137	8202	
cholesterol metabolic process	0.008	0.021	0.084	0.117	0.053	0.042	0.928	0.285	0.149	8203	
blood pressure regulation	0.599	0.968	0.124	0.156	0.214	0.264	0.445	0.718	0.665	8217	
cell death	0.036	0.769	0.612	0.187	0.166	0.047	0.772	0.650	0.416	8219	
sulfate transport	0.447	0.637	0.897	0.244	0.532	0.934	0.303	0.489	0.408	8272	
regulation of G-protein coupled receptor protein signaling pathway	0.242	0.552	0.353	0.940	0.244	0.055	0.584	0.142	0.874	8277	
cell proliferation	0.321	0.007	0.766	0.018	0.086	0.010	0.318	0.173	0.280	8283	
positive regulation of cell proliferation	0.303	0.139	0.861	0.110	0.164	0.096	0.325	0.084	0.194	8284	
negative regulation of cell proliferation	0.045	0.063	0.556	0.214	0.222	0.080	0.055	0.095	0.090	8285	
insulin receptor signaling pathway	0.130	0.179	0.017	0.085	0.654	0.216	0.039	0.279	0.054	8286	
isoprenoid biosynthetic process	0.852	0.406	0.222	0.286	0.052	0.433	0.310	0.417	0.070	8299	
adult locomotor behavior	0.934	0.262	0.136	0.414	0.578	0.097	0.006	0.267	0.258	8344	
regulation of cell shape	0.043	0.010	0.126	0.026	0.033	0.018	0.251	0.006	0.120	8360	
RNA splicing	0.694	0.010	0.012	0.018	0.017	0.005	0.346	0.070	0.016	8380	
fibroblast growth factor receptor signaling pathway	0.139	0.195	0.389	0.928	0.410	0.196	0.090	0.166	0.682	8543	
epidermis development	0.879	0.968	0.960	0.963	0.679	0.822	0.683	0.995	0.978	8544	
male gonad development	0.844	0.919	0.972	0.771	0.833	0.752	0.402	0.282	0.812	8584	
lipid biosynthetic process	0.157	0.194	0.672	0.024	0.003	0.035	0.057	0.471	0.291	8610	
induction of apoptosis by extracellular signals	0.078	0.410	0.578	0.008	0.251	0.857	0.455	0.297	0.878	8624	
induction of apoptosis via death domain receptors	0.972	0.419	0.535	0.793	0.429	0.540	0.909	0.292	0.940	8625	
induction of apoptosis by intracellular signals	0.573	0.105	0.115	0.155	0.410	0.769	0.781	0.127	0.150	8629	
apoptotic program	0.404	0.342	0.243	0.219	0.445	0.248	0.374	0.030	0.033	8632	
carbohydrate transport	0.513	0.567	0.341	0.656	0.721	0.627	0.273	0.945	0.771	8643	
amino acid biosynthetic process	0.287	0.036	0.030	0.492	0.337	0.449	0.748	0.690	0.622	8652	
phospholipid biosynthetic process	0.176	0.537	0.069	0.203	0.101	0.232	0.573	0.838	0.224	8654	
biosynthetic process	0.931	0.040	0.100	0.224	0.015	0.125	0.560	0.278	0.178	9058	
aromatic amino acid family metabolic process	0.845	0.545	0.981	0.660	0.395	0.468	0.466	0.366	0.849	9072	
nucleoside metabolic process	0.815	0.366	0.575	0.381	0.341	0.936	0.233	0.426	0.492	9116	
nucleotide metabolic process	0.939	0.172	0.039	0.063	0.051	0.052	0.157	0.368	0.114	9117	
nucleotide biosynthetic process	0.390	0.656	0.250	0.292	0.689	0.393	0.295	0.949	0.847	9165	
cyclic nucleotide biosynthetic process	0.429	0.466	0.660	0.079	0.162	0.248	0.710	0.209	0.341	9190	
protein secretion	0.324	0.899	0.851	0.062	0.103	0.015	0.705	0.566	0.571	9306	
oligosaccharide biosynthetic process	0.127	0.296	0.030	0.401	0.727	0.648	0.094	0.342	0.495	9312	
response to radiation	0.815	0.234	0.962	0.162	0.374	0.205	0.020	0.696	0.284	9314	
phosphoenolpyruvate-dependent sugar phosphotransferase sys	0.331	0.985	0.434	0.266	0.036	0.293	0.380	0.676	0.243	9401	
response to heat	0.710	0.227	0.672	0.364	0.110	0.095	0.560	0.068	0.662	9408	
response to cold	0.662	0.107	0.256	0.248	0.036	0.191	0.576	0.760	0.790	9409	
response to UV	0.107	0.143	0.642	0.261	0.812	0.205	0.689	0.112	0.181	9411	
response to biotic stimulus	0.068	0.448	0.979	0.401	1.000	0.672	0.729	0.821	0.075	9607	
response to wounding	0.134	0.116	0.802	0.639	0.095	0.415	0.336	0.292	0.788	9611	
response to virus	0.888	0.958	0.669	0.935	0.908	0.863	0.934	0.919	0.788	9615	
response to toxin	0.123	0.305	0.823	0.135	0.178	0.015	0.523	0.421	0.556	9636	
anatomical structure morphogenesis	0.305	0.437	0.637	0.502	0.503	0.326	0.538	0.564	0.973	9653	
hormone-mediated signaling	0.832	0.590	0.852	0.571	0.421	0.150	0.726	0.309	0.552	9755	
embryonic development	0.770	0.606	0.569	0.043	0.205	0.094	0.592	0.131	0.071	9790	
organ morphogenesis	0.714	0.553	0.853	0.900	0.063	0.273	0.702	0.336	0.948	9887	
tissue development	0.880	0.135	0.983	0.643	0.451	0.201	0.879	0.218	0.358	9888	
anterior/posterior pattern formation	0.496	0.729	0.113	0.295	0.219	0.408	0.631	0.429	0.336	9952	
dorsal/ventral pattern formation	0.503	0.205	0.884	0.870	0.503	0.376	0.145	0.526	0.449	10001	
regulation of signal transduction	0.219	0.597	0.328	0.339	0.182	0.446	0.361	0.253	0.421	9966	
negative regulation of signal transduction	0.219	0.597	0.328	0.339	0.182	0.446	0.361	0.253	0.421	9966	
glial cell differentiation	0.274	0.205	0.884	0.870	0.503	0.376	0.145	0.526	0.449	10001	
gastrulation (sensu Mammalia)	0.547	0.081	0.037	0.374	0.063	0.022	0.078	0.455	0.824	10003	
protein transport	0.100	0.008	0.005	0.008	0.007	0.005	0.350	0.060	0.016	15031	
DNA integration	0.333	0.510	0.830	0.414	0.140	0.290	0.868	0.199	0.184	15074	
oxygen transport	0.978	0.068	0.140	0.654	0.830	0.945	0.342	0.371	0.070	15671	
organic anion transport	0.874	0.972	0.772	0.699	0.640	0.911	0.378	0.930	0.381	15711	
glucose transport	0.226	0.317</									

Supplemental Table 1. Complete list of Gene Ontology Biological Processes from the SAFE analysis												
GO term	Neocortex p-values			Heart p-values			Muscle p-values			GO ID	SAFE	
	Age	CR	Resv	Age	CR	Resv	Age	CR	Resv			
cell growth	0.615	0.655	0.618	0.035	0.016	0.023	0.800	0.166	0.347	16049		
carbohydrate biosynthetic process	0.305	0.260	0.550	0.144	0.647	0.290	0.923	0.006	0.520	16051		
Wnt receptor signaling pathway	0.959	0.016	0.198	0.418	0.157	0.398	0.472	0.336	0.453	16055		
immunoglobulin mediated immune response	0.107	0.911	0.960	0.492	0.561	0.877	0.736	0.241	0.105	16064		
mRNA metabolic process	0.775	0.055	0.581	0.162	0.016	0.026	0.398	0.212	0.036	16071		
sterol biosynthetic process	0.169	0.356	0.866	0.014	0.044	0.248	0.060	0.088	0.292	16126		
snRNA processing	0.089	0.755	0.244	0.771	0.166	0.951	0.443	0.511	0.151	16180		
vesicle-mediated transport	0.122	0.018	0.006	0.074	0.186	0.083	0.040	0.272	0.125	16192		
endosome transport	0.223	0.658	0.631	0.068	0.235	0.085	0.481	0.097	0.203	16197		
phosphorylation	0.153	0.188	0.006	0.382	0.257	0.429	0.020	0.084	0.626	16310		
morphogenesis of embryonic epithelium	0.189	0.132	0.150	0.481	0.226	0.752	0.211	0.705	0.963	16331		
cell-cell adhesion	0.029	0.215	0.157	0.188	0.542	0.371	0.146	0.523	0.520	16337		
calcium-independent cell-cell adhesion	0.497	0.950	0.969	0.753	0.992	0.923	0.673	0.633	0.718	16338		
calcium-dependent cell-cell adhesion	0.822	0.856	0.225	0.853	0.554	0.269	0.865	0.380	0.900	16339		
dendrite development	0.060	0.774	0.855	0.280	0.630	0.560	0.042	0.274	0.519	16358		
cell migration	0.403	0.202	0.025	0.064	0.007	0.005	0.412	0.538	0.409	16477		
negative regulation of transcription	0.040	0.338	0.048	0.081	0.074	0.032	0.753	0.622	0.355	16481		
protein processing	0.119	0.229	0.045	0.733	0.671	0.368	0.043	0.138	0.115	16485		
negative regulation of angiogenesis	0.813	0.461	0.419	0.281	0.210	0.524	0.807	0.263	0.044	16525		
protein ubiquitination	0.368	0.101	0.731	0.305	0.015	0.046	0.212	0.358	0.512	16567		
chromatin modification	0.635	0.023	0.005	0.060	0.028	0.015	0.343	0.281	0.012	16568		
histone methylation	0.488	0.137	0.114	0.943	0.230	0.385	0.654	0.190	0.029	16571		
cell wall catabolic process	0.803	0.680	0.738	0.123	0.444	0.130	0.391	0.954	0.863	16998		
peptidyl-serine phosphorylation	0.104	0.427	0.310	0.656	0.509	0.192	0.968	0.314	0.198	18105		
peptidyl-tyrosine phosphorylation	0.061	0.165	0.780	0.086	0.106	0.253	0.212	0.254	0.298	18108		
peptide cross-linking	0.122	0.852	0.906	0.592	0.953	0.715	0.042	0.540	0.903	18149		
protein amino acid N-linked glycosylation via asparagine	0.172	0.238	0.006	0.008	0.188	0.107	0.840	0.379	0.131	18279		
protein-chromophore linkage	0.180	0.414	0.511	0.395	0.752	0.965	0.261	0.847	0.474	18298		
protein amino acid prenylation	0.475	0.420	0.274	0.292	0.236	0.239	0.542	0.861	0.666	18346		
cytokine and chemokine mediated signaling pathway	0.840	0.182	0.955	0.128	0.036	0.147	0.279	0.189	0.302	19221		
sensory perception of pain	0.071	0.032	0.187	0.353	0.111	0.358	0.618	0.814	0.739	19233		
response to pheromone	0.863	0.914	0.992	0.930	0.736	0.870	0.397	0.055	0.407	19236		
pyridine nucleotide biosynthetic process	0.858	0.599	0.456	0.691	0.141	0.434	0.936	0.488	0.771	19363		
leukotriene biosynthetic process	0.664	0.748	0.636	0.087	0.608	0.157	0.969	0.465	0.265	19370		
protein metabolic process	0.299	0.042	0.006	0.718	0.045	0.065	0.367	0.368	0.670	19538		
calcium-mediated signaling	0.198	0.988	0.862	0.807	0.327	0.647	0.419	0.264	0.671	19722		
antimicrobial humoral response (sensu Vertebrata)	0.106	0.909	0.787	0.513	0.888	0.988	0.077	0.441	0.903	19735		
cytolysis	0.598	0.748	0.603	0.424	0.940	0.598	0.720	0.959	0.866	19835		
antigen processing and presentation	0.247	0.927	0.664	0.925	0.627	0.891	0.447	0.808	0.564	19882		
antigen processing and presentation of exogenous peptide antigen	0.023	0.411	0.714	0.569	0.469	0.810	0.577	0.428	0.160	19886		
metal ion transport	0.856	0.147	0.221	0.252	0.379	0.717	0.820	0.962	0.977	30001		
lamellipodium biogenesis	0.101	0.432	0.332	0.008	0.031	0.182	0.333	0.191	0.402	30032		
actin cytoskeleton organization and biogenesis	0.070	0.003	0.015	0.008	0.016	0.005	0.186	0.053	0.226	30036		
actin filament polymerization	0.471	0.290	0.015	0.065	0.589	0.257	0.044	0.625	0.776	30041		
actin filament-based movement	0.668	0.437	0.538	0.472	0.125	0.341	0.012	0.252	0.025	30048		
insulin secretion	0.668	0.321	0.032	0.926	0.796	0.795	0.425	0.205	0.809	30073		
hemopoiesis	0.311	0.825	0.460	0.212	0.007	0.445	0.296	0.368	0.207	30097		
myeloid cell differentiation	0.400	0.077	0.147	0.175	0.051	0.320	0.290	0.839	0.872	30099		
regulation of endocytosis	0.138	0.095	0.006	0.283	0.090	0.141	0.340	0.211	0.360	30100		
cell differentiation	0.087	0.064	0.646	0.111	0.220	0.298	0.020	0.311	0.097	30154		
regulation of cell adhesion	0.104	0.024	0.067	0.547	0.033	0.044	0.283	0.429	0.518	30155		
protein catabolic process	0.188	0.032	0.053	0.028	0.023	0.005	0.076	0.399	0.293	30163		
platelet activation	0.796	0.937	0.618	0.752	0.075	0.238	0.169	0.440	0.913	30168		
negative regulation of Wnt receptor signaling pathway	0.561	0.120	0.878	0.239	0.434	0.603	0.751	0.367	0.297	30178		
neuron differentiation	0.780	0.841	0.597	0.291	0.760	0.675	0.257	0.906	0.897	30182		
B cell differentiation	0.676	0.946	0.627	0.655	0.405	0.259	0.643	0.337	0.470	30183		
extracellular matrix organization and biogenesis	0.369	0.148	0.684	0.064	0.162	0.414	0.957	0.432	0.580	30198		
collagen fibril organization	0.704	0.329	0.052	0.576	0.124	0.186	0.275	0.891	0.870	30199		
keratinocyte differentiation	0.133	0.714	0.930	0.820	0.671	0.973	0.968	0.372	0.701	30216		
T cell differentiation	0.217	0.225	0.056	0.936	0.732	0.729	0.046	0.926	0.803	30217		
erythrocyte differentiation	0.348	0.133	0.542	0.168	0.953	0.537	0.561	0.932	0.134	30218		
bone mineralization	0.143	0.787	0.618	0.924	0.822	0.671	0.192	0.021	0.398	30282		
negative regulation of cell growth	0.106	0.041	0.102	0.330	0.116	0.717	0.104	0.104	0.499	30308		
sperm motility	0.869	0.783	0.302	0.234	0.480	0.650	0.189	0.528	0.814	30317		
melanocyte differentiation	0.669	0.152	0.054	0.901	0.433	0.553	0.742	0.006	0.162	30318		
lung development	0.920	0.144	0.497	0.306	0.089	0.095	0.716	0.157	0.894	30324		
embryonic limb morphogenesis	0.333	0.138	0.793	0.774	0.741	0.528	0.429	0.023	0.639	30326		
regulation of cell migration	0.227	0.010	0.481	0.177	0.015	0.377	0.585	0.809	0.375	30334		
positive regulation of cell migration	0.330	0.259	0.607	0.036	0.731	0.104	0.436	0.616	0.342	30335		
negative regulation of cell migration	0.504	0.105	0.154	0.108	0.048	0.688	0.283	0.402	0.067	30336		
ER-associated protein catabolic process	0.711	0.652	0.364	0.588	0.109	0.368	0.249	0.144	0.272	30433		
BMP signaling pathway	0.049	0.332	0.101	0.235	0.228	0.049	0.092	0.029	0.032	30509		
negative regulation of BMP signaling pathway	0.868	0.249	0.313	0.441	0.007	0.052	0.157	0.416	0.770	30514		
estrogen receptor signaling pathway	0.575	0.091	0.615	0.028	0.126	0.005	0.016	0.718	0.923	30520		
androgen receptor signaling pathway	0.668	0.020	0.062	0.099	0.059	0.005	0.224	0.359	0.255	30521		
collagen catabolic process	0.219	0.819	0.968	0.950	0.734	0.689	0.180	0.719	0.954	30574		
neutrophil chemotaxis	0.114	0.597	0.583	0.073	0.752	0.216	0.598	0.414	0.772	30593		
epithelial cell differentiation	0.663	0.953	0.956	0.820	0.965	0.946	0.893	0.148	0.759	30855		
cortical actin cytoskeleton organization and biogenesis	0.008	0.048	0.062	0.131	0.023	0.211	0.357	0.032	0.031	30866		
mammary gland development	0.951	0.077	0.143	0.386	0.931	0.914	0.110	0.765	0.730	30879		
positive regulation of B cell proliferation	0.135	0.887	0.861	0.558	0.680	0.857	0.171	0.114	0.169	30890		
forebrain development	0.091	0.042	0.108	0.716	0.568	0.325	0.380	0.066	0.189	30900		
midbrain development	0.676	0.795	0.906	0.094	0.684	0.739	0.504	0.757	0.567	30901		
hindbrain development	0.392	0.701	0.655	0.345	0.781	0.622	0.461	0.727	0.390	30902		
unfolded protein response	0.202	0.867	0.555	0.038	0.808	0.716	0.490	0.558	0.552	30968		
hair follicle morphogenesis	0.746	0.507	0.559	0.586	0.419	0.046	0.158	0.314	0.632	31069		
neurite development	0.281	0.311	0.352	0.285	0.598	0.202	0.079	0.826	0.793	31175		
keratinization	0.764	0.980	0.935	1.000	0.890	0.982	0.948	0.843	0.962	31424		
regulation of ARF protein signal transduction	0.194	0.054	0.103	0.104	0.152	0.662	0.791	0.221	0.071	32012		
regulation of Rho protein signal transduction	0.144	0.018	0.044	0.087	0.427	0.161	0.716	0.807	0.948			

Supplemental Table 1. Complete list of Gene Ontology Biological Processes from the SAFE analysis												
GO term	Neocortex p-values			Heart p-values			Muscle p-values			GO ID	SAFE	
	Age	CR	Resv	Age	CR	Resv	Age	CR	Resv			
B cell activation	0.906	0.544	0.600	0.984	0.333	0.283	0.572	0.164	0.720	42113		
regulation of cell proliferation	0.781	0.020	0.024	0.086	0.036	0.142	0.006	0.012	0.076	42127		
negative regulation of T cell proliferation	0.328	0.912	0.581	0.157	0.273	0.630	0.032	0.517	0.722	42130		
lipoprotein metabolic process	0.016	0.992	0.632	0.728	0.056	0.453	0.927	0.723	0.800	42157		
ribosome biogenesis and assembly	0.398	0.087	0.796	0.078	0.527	0.689	0.017	0.378	0.179	42254		
inner ear morphogenesis	0.711	0.580	0.726	0.611	0.732	0.960	0.808	0.811	0.681	42472		
odontogenesis (sensu Vertebrata)	0.296	0.217	0.912	0.529	0.339	0.254	0.705	0.497	0.667	42475		
auditory receptor cell differentiation	0.116	0.122	0.047	0.918	0.540	0.924	0.821	0.725	0.046	42491		
response to drug	0.530	0.614	0.365	0.734	0.345	0.227	0.012	0.139	0.086	42493		
myelination	0.146	0.192	0.020	0.559	0.092	0.164	0.745	0.945	0.243	42552		
glucose homeostasis	0.618	0.258	0.054	0.123	0.167	0.228	0.659	0.143	0.748	42593		
cholesterol homeostasis	0.109	0.207	0.254	0.289	0.034	0.010	0.134	0.693	0.321	42632		
embryonic digit morphogenesis	0.317	0.182	0.727	0.058	0.193	0.295	0.312	0.138	0.599	42733		
defense response to bacterium	0.437	1.000	1.000	0.909	0.890	0.912	0.767	0.724	0.988	42742		
regulation of apoptosis	0.188	0.244	0.006	0.019	0.015	0.013	0.312	0.216	0.338	42981		
positive regulation of apoptosis	0.034	0.400	0.852	0.008	0.024	0.042	0.148	0.209	0.036	43065		
negative regulation of apoptosis	0.549	0.091	0.082	0.347	0.007	0.043	0.037	0.036	0.048	43066		
regulation of GTPase activity	0.320	0.023	0.090	0.255	0.722	0.094	0.213	0.369	0.334	43087		
positive regulation of I-kappaB kinase/NF-kappaB cascade	0.582	0.534	0.457	0.668	0.047	0.053	0.889	0.099	0.475	43123		
pigmentation	0.749	0.306	0.197	0.866	0.455	0.326	0.885	0.013	0.100	43473		
negative regulation of neuron apoptosis	0.025	0.930	0.407	0.445	0.083	0.139	0.082	0.541	0.266	43524		
cellular metabolic process	0.351	0.040	0.040	0.120	0.439	0.259	0.567	0.678	0.181	44237		
cellular protein metabolic process	0.453	0.132	0.074	0.149	0.163	0.603	0.006	0.027	0.075	44267		
positive regulation of interferon-gamma biosynthetic process	0.896	0.850	0.907	0.529	0.920	0.826	0.123	0.505	0.397	45078		
positive regulation of interleukin-2 biosynthetic process	0.394	0.182	0.285	0.682	0.909	0.837	0.683	0.376	0.046	45086		
innate immune response	0.179	0.629	0.946	0.790	0.634	0.244	0.433	0.734	0.791	45087		
cell fate commitment	0.256	0.695	0.695	0.324	0.979	0.920	0.825	0.276	0.743	45165		
fat cell differentiation	0.092	0.123	0.110	0.017	0.606	0.935	0.427	0.728	0.107	45444		
regulation of transcription	0.286	0.912	0.750	0.579	0.264	0.292	0.560	0.376	0.688	45449		
bone resorption	0.702	0.266	0.579	0.638	0.641	0.389	0.182	0.219	0.634	45453		
cell redox homeostasis	0.235	0.319	0.464	0.441	0.091	0.178	0.612	0.218	0.041	45454		
regulation of cell differentiation	0.849	0.024	0.006	0.040	0.389	0.160	0.505	0.404	0.738	45595		
negative regulation of cell differentiation	0.198	0.111	0.158	0.186	0.041	0.400	0.596	0.059	0.579	45596		
regulation of neuron differentiation	0.122	0.474	0.225	0.071	0.560	0.314	0.807	0.499	0.390	45664		
negative regulation of neuron differentiation	0.641	0.376	0.304	0.459	0.240	0.164	0.342	0.750	0.902	45665		
negative regulation of progression through cell cycle	0.036	0.020	0.049	0.597	0.069	0.034	0.264	0.187	0.028	45786		
positive regulation of protein kinase activity	0.174	0.180	0.844	0.798	0.741	0.204	0.091	0.701	0.335	45860		
negative regulation of transcription, DNA-dependent	0.475	0.018	0.015	0.717	0.013	0.036	0.313	0.069	0.018	45892		
positive regulation of transcription, DNA-dependent	0.008	0.045	0.015	0.237	0.039	0.025	0.496	0.156	0.046	45893		
positive regulation of transcription	0.265	0.017	0.220	0.312	0.345	0.103	0.069	0.025	0.079	45941		
positive regulation of transcription from RNA polymerase II prom	0.459	0.094	0.637	0.033	0.176	0.262	0.172	0.008	0.003	45944		
positive regulation of translation	0.932	0.010	0.026	0.008	0.033	0.013	0.552	0.706	0.125	45946		
positive regulation of JNK cascade	0.988	0.743	0.488	0.327	0.245	0.185	0.130	0.153	0.134	46330		
protein amino acid autophosphorylation	0.053	0.132	0.961	0.527	0.179	0.514	0.404	0.043	0.016	46777		
phospholipid dep phosphorylation	0.355	0.027	0.050	0.040	0.064	0.091	0.054	0.068	0.135	46839		
intracellular transport	0.067	0.021	0.026	0.513	0.099	0.393	0.780	0.014	0.003	46907		
phosphoinositide-mediated signaling	0.317	0.774	0.802	0.204	0.410	0.153	0.670	0.975	0.788	48015		
focal adhesion formation	0.471	0.010	0.026	0.073	0.166	0.152	0.398	0.640	0.534	48041		
pigmentation during development	0.785	0.770	0.170	0.454	0.047	0.012	0.409	0.070	0.236	48066		
regulation of synaptic plasticity	0.188	0.087	0.036	0.789	0.386	0.718	0.014	0.084	0.497	48167		
cell development	0.236	0.178	0.595	0.406	0.250	0.092	0.768	0.884	0.636	48468		
cell maturation	0.473	0.089	0.101	0.805	0.388	0.915	0.070	0.800	0.994	48469		
rhythmic process	0.021	0.408	0.343	0.143	0.261	0.906	0.854	0.048	0.326	48511		
blood vessel morphogenesis	0.877	0.169	0.150	0.250	0.142	0.096	0.634	0.788	0.225	48514		
lymph node development	0.606	0.533	0.870	0.917	0.196	0.041	0.086	0.042	0.859	48535		
thymus development	0.937	0.638	0.830	0.790	0.925	0.525	0.663	0.379	0.346	48538		
morphogenesis (sensu Mammalia)	0.434	0.179	0.089	0.230	0.007	0.017	0.827	0.529	0.663	48595		
neuron fate commitment	0.170	0.123	0.622	0.220	0.475	0.618	0.846	0.722	0.809	48663		
neuron development	0.859	0.949	0.983	0.262	0.720	0.158	0.523	0.561	0.967	48666		
branching morphogenesis of a tube	0.596	0.136	0.470	0.671	0.353	0.528	0.642	0.182	0.802	48754		
positive regulation of epithelial cell proliferation	0.224	0.540	0.569	0.199	0.466	0.117	0.625	0.385	0.190	50679		
positive regulation of peptidyl-tyrosine phosphorylation	0.238	0.789	0.914	0.376	0.362	0.998	0.356	0.794	0.229	50731		
positive regulation of phagocytosis	0.038	0.462	0.829	0.147	0.057	0.331	0.008	0.037	0.064	50766		
regulation of immune response	0.403	0.944	0.568	0.812	0.413	0.393	0.347	0.031	0.302	50776		
synapse organization and biogenesis	0.938	0.141	0.318	0.770	0.618	0.329	0.091	0.790	0.609	50808		
protein stabilization	0.338	0.372	0.932	0.493	0.377	0.632	0.027	0.700	0.546	50821		
T cell receptor signaling pathway	0.468	0.397	0.181	0.763	0.468	0.799	0.277	0.793	0.778	50852		
cellular physiological process	0.305	0.020	0.121	0.026	0.169	0.023	0.033	0.137	0.601	50875		
regulation of blood vessel size	0.570	0.975	0.568	0.281	0.751	0.813	0.704	0.500	0.378	50880		
regulation of balance	0.422	0.702	0.043	0.019	0.860	0.170	0.070	0.475	0.077	50885		
response to stimulus	0.878	0.724	0.924	0.723	0.964	0.975	0.818	0.847	0.990	50896		
barbed-end actin filament capping	0.591	0.435	0.701	0.023	0.071	0.377	0.615	0.612	0.325	51016		
actin filament bundle formation	0.302	0.162	0.599	0.244	0.167	0.195	0.149	0.444	0.120	51017		
mRNA transport	0.644	0.204	0.062	0.416	0.496	0.005	0.320	0.141	0.038	51028		
regulation of small GTPase mediated signal transduction	0.022	0.010	0.034	0.168	0.072	0.098	0.455	0.108	0.278	51056		
chaperone cofactor-dependent protein folding	0.599	0.296	0.420	0.572	0.748	0.993	0.902	0.781	0.191	51085		
activation of NF-kappaB transcription factor	0.443	0.223	0.914	0.863	0.718	0.106	0.463	0.053	0.155	51092		
cartilage development	0.697	0.461	0.470	0.182	0.884	0.038	0.639	0.585	0.285	51216		
protein polymerization	0.908	0.809	0.154	0.121	0.690	0.045	0.358	0.730	0.966	51258		
protein homooligomerization	0.872	0.724	0.808	0.412	0.051	0.102	0.103	0.324	0.295	51260		
cell division	0.469	0.099	0.006	0.312	0.007	0.005	0.297	0.656	0.267	51301		